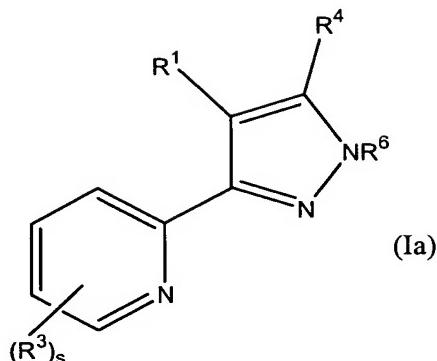


The claimed invention is:

1. A compound of formula (Ia):



or a pharmaceutically acceptable salt, prodrug, tautomer, hydrate or solvate thereof,

5 wherein:

- R¹ is a saturated, unsaturated, or aromatic C₃-C₂₀ mono-, bi- or polycyclic ring optionally containing at least one heteroatom selected from the group consisting of N, O and S, wherein R¹ can optionally be further independently substituted with at least one moiety independently selected from the group consisting of: carbonyl,
10 halo, halo(C₁-C₆)alkyl, perhalo(C₁-C₆)alkyl, perhalo(C₁-C₆)alkoxy,
(C₁-C₆)alkyl, (C₂-C₆)alkenyl, (C₂-C₆)alkynyl, hydroxy, oxo, mercapto, (C₁-C₆)alkylthio, (C₁-C₆)alkoxy, (C₅-C₁₀)aryl or (C₅-C₁₀)heteroaryl, (C₅-C₁₀)aryloxy or
(C₅-C₁₀)heteroaryloxy, (C₅-C₁₀)ar(C₁-C₆)alkyl or (C₅-C₁₀)heteroar(C₁-C₆)alkyl,
(C₅-C₁₀)ar(C₁-C₆)alkoxy or (C₅-C₁₀)heteroar(C₁-C₆)alkoxy, HO-(C=O)-, ester, amido,
15 ether, amino, amino(C₁-C₆)alkyl, (C₁-C₆)alkylamino(C₁-C₆)alkyl,
di(C₁-C₆)alkylamino(C₁-C₆)alkyl, (C₅-C₁₀)heterocycl(C₁-C₆)alkyl, (C₁-C₆)alkyl- and
di(C₁-C₆)alkylamino, cyano, nitro, carbamoyl, (C₁-C₆)alkylcarbonyl,
(C₁-C₆)alkoxycarbonyl, (C₁-C₆)alkylaminocarbonyl,
di(C₁-C₆)alkylaminocarbonyl, (C₅-C₁₀)arylcarbonyl, (C₅-C₁₀)aryloxycarbonyl,
20 (C₁-C₆)alkylsulfonyl, and (C₅-C₁₀)arylsulfonyl;

each R³ is independently selected from the group consisting of: hydrogen,
halo, halo(C₁-C₆)alkyl, (C₁-C₆)alkyl, (C₂-C₆)alkenyl, (C₂-C₆)alkynyl,
perhalo(C₁-C₆)alkyl, phenyl, (C₅-C₁₀)heteroaryl, (C₅-C₁₀)heterocyclic,

(C₃-C₁₀)cycloalkyl, hydroxy, (C₁-C₆)alkoxy, perhalo(C₁-C₆)alkoxy, phenoxy,
(C₅-C₁₀)heteroaryl-O-, (C₅-C₁₀)heterocyclic-O-, (C₃-C₁₀)cycloalkyl-O-,
(C₁-C₆)alkyl-S-, (C₁-C₆)alkyl-SO₂-, (C₁-C₆)alkyl-NH-SO₂-, O₂N-, NC-, amino,
Ph(CH₂)₁₋₆HN-, (C₁-C₆)alkyl HN-, (C₁-C₆)alkylamino, [(C₁-C₆)alkyl]₂-amino,
5 (C₁-C₆)alkyl-SO₂-NH-, amino(C=O)-, aminoO₂S-, (C₁-C₆)alkyl-(C=O)-NH-,
(C₁-C₆)alkyl-(C=O)-[((C₁-C₆)alkyl)-N]-, phenyl-(C=O)-NH-,
phenyl-(C=O)-[(C₁-C₆)alkyl)-N]-, (C₁-C₆)alkyl-(C=O)-, phenyl-(C=O)-,
(C₅-C₁₀)heteroaryl-(C=O)-, (C₅-C₁₀)heterocyclic-(C=O)-, (C₃-C₁₀)cycloalkyl-(C=O)-,
HO-(C=O)-, (C₁-C₆)alkyl-O-(C=O)-, H₂N(C=O)-, (C₁-C₆)alkyl-NH-(C=O)-,
10 [(C₁-C₆)alkyl]₂-N-(C=O)-, phenyl-NH-(C=O)-, phenyl-[(C₁-C₆)alkyl)-N]--(C=O)-,
(C₅-C₁₀)heteroaryl-NH-(C=O)-, (C₅-C₁₀)heterocyclic-NH-(C=O)-,
(C₃-C₁₀)cycloalkyl-NH-(C=O)- and (C₁-C₆)alkyl-(C=O)-O-;

where alkyl, alkenyl, alkynyl, phenyl, heteroaryl, heterocyclic, cycloalkyl,
15 alkoxy, phenoxy, amino of R³ is optionally substituted by at least one substituent
independently selected from (C₁-C₆)alkyl, (C₁-C₆)alkoxy, halo(C₁-C₆)alkyl, halo,
H₂N-, Ph(CH₂)₁₋₆HN-, and (C₁-C₆)alkylHN-;

s is an integer from one to five;

20 R⁴ is selected from the group consisting of: hydrogen, halo, halo(C₁-C₆)alkyl, (C₁-C₆)alkyl, (C₂-C₆)alkenyl, (C₂-C₆)alkynyl,
perhalo(C₁-C₆)alkyl, phenyl, (C₅-C₁₀)heteroaryl, (C₅-C₁₀)heterocyclic,
(C₃-C₁₀)cycloalkyl, hydroxy, (C₁-C₆)alkoxy, perhalo(C₁-C₆)alkoxy, phenoxy,
25 (C₅-C₁₀)heteroaryl-O-, (C₅-C₁₀)heterocyclic-O-, (C₃-C₁₀)cycloalkyl-O-,
(C₁-C₆)alkyl-S-, (C₁-C₆)alkyl-SO₂-, (C₁-C₆)alkyl-NH-SO₂-, O₂N-, NC-, amino,
Ph(CH₂)₁₋₆NH-, alkylNH-, (C₁-C₆)alkylamino, [(C₁-C₆)alkyl]₂-amino,
(C₁-C₆)alkyl-SO₂-NH-, amino(C=O)-, aminoSO₂-, (C₁-C₆)alkyl-(C=O)-NH-,
(C₁-C₆)alkyl-(C=O)-((C₁-C₆)alkyl)-N]-, phenyl-(C=O)-NH-,
30 phenyl-(C=O)-((C₁-C₆)alkyl)-N]-, (C₁-C₆)alkyl-(C=O)-, phenyl-(C=O)-,
(C₅-C₁₀)heteroaryl-(C=O)-, (C₅-C₁₀)heterocyclic-(C=O)-, cycloalkyl-(C=O)-,

HO-(C=O)-, (C₁-C₆)alkyl-O-(C=O)-, H₂N(C=O)-, (C₁-C₆)alkyl-NH-(C=O)-,
((C₁-C₆)alkyl)₂-N-(C=O)-, phenyl-NH-(C=O)-, phenyl-((C₁-C₆)alkyl)-N]--(C=O)-,
(C₅-C₁₀)heteroaryl-NH-(C=O)-, (C₅-C₁₀)heterocyclic-NH-(C=O)-,
(C₃-C₁₀)cycloalkyl-NH-(C=O)- and (C₁-C₆)alkyl-(C=O)-O-,

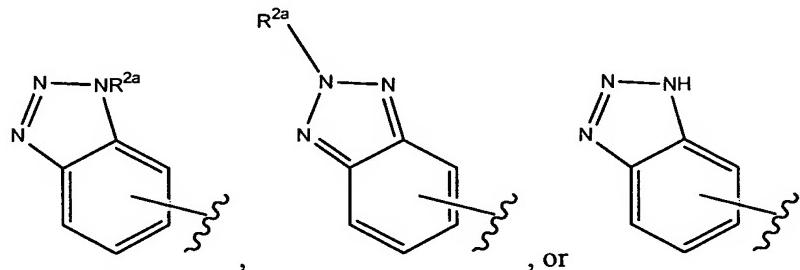
5 where alkyl, alkenyl, alkynyl, phenyl, heteroaryl, heterocyclic, cycloalkyl,
alkoxy, phenoxy, and amino of R⁴ is optionally substituted by at least one
substituent independently selected from the group consisting of (C₁-C₆)alkyl, (C₁-
C₆)alkoxy, halo(C₁-C₆)alkyl, halo, H₂N-, Ph(CH₂)₁₋₆-NH-, and (C₁-C₆)alkylNH-; and

10 R⁶ is selected from the group consisting of hydrogen, (C₁-C₆)alkyl,
(C₂-C₆)alkenyl, (C₂-C₆)alkynyl, phenyl, (C₅-C₁₀)heteroaryl, (C₅-C₁₀)heterocyclic,
(C₃-C₁₀)cycloalkyl, (C₁-C₆)alkyl-(SO₂)-, phenyl-(SO₂)-, H₂N-(SO₂)-,
(C₁-C₆)alkyl-NH-(SO₂)-, ((C₁-C₆)alkyl)₂N-(SO₂)-, phenyl-NH-(SO₂)-,
(phenyl)₂N-(SO₂)-, (C₁-C₆)alkyl-(C=O)-, phenyl-(C=O)-, (C₅-C₁₀)heteroaryl-(C=O)-,
15 (C₅-C₁₀)heterocyclic-(C=O)-, (C₃-C₁₀)cycloalkyl-(C=O)-, (C₁-C₆)alkyl-O-(C=O)-,
(C₅-C₁₀)heterocyclic-O-(C=O)-, (C₃-C₁₀)cycloalkyl-O-(C=O)-, H₂N-(C=O)-,
(C₁-C₆)alkyl-NH-(C=O)-, phenyl-NH-(C=O)-, (C₅-C₁₀)heteroaryl-NH-(C=O)-,
(C₅-C₁₀)heterocyclic-NH-(C=O)-, (C₃-C₁₀)cycloalkyl-NH-(C=O)-,
15 ((C₁-C₆)alkyl)₂N-(C=O)-, (phenyl)₂N-(C=O)-, phenyl-[((C₁-C₆)alkyl)-N]--(C=O)-,
20 (C₅-C₁₀)heteroaryl-[((C₁-C₆)alkyl)-N]--(C=O)-,
(C₅-C₁₀)heterocyclic-[((C₁-C₆)alkyl)-N]--(C=O)-, and
(C₃-C₁₀)cycloalkyl-[((C₁-C₆)alkyl)-N]--(C=O)-;

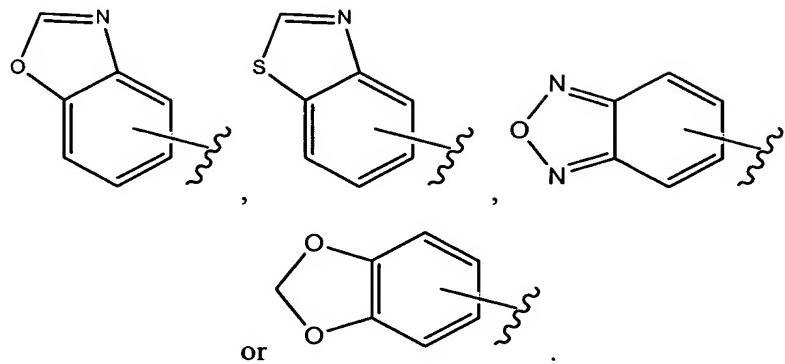
where alkyl, alkenyl, alkynyl, phenyl, benzyl, heteroaryl, heterocyclic,
cycloalkyl, alkoxy, phenoxy, amino of R⁶ is optionally substituted with at least one
25 moiety independently selected from the group consisting of halo, (C₁-C₆)alkyl,
(C₂-C₆)alkenyl, (C₂-C₆)alkynyl, perhalo(C₁-C₆)alkyl, (C₃-C₁₀)cycloalkyl, phenyl,
benzyl, (C₅-C₁₀)heterocyclic, (C₅-C₁₀)heteroaryl, (C₁-C₆)alkyl-SO₂-, formyl, NC-,
(C₁-C₆)alkyl-(C=O)-, (C₃C₁₀)cycloalkyl-(C=O)-, phenyl-(C=O)-,
(C₅-C₁₀)heterocyclic-(C=O)-, (C₅-C₁₀)heteroaryl-(C=O)-, HO-(C=O)-,
30 (C₁-C₆)alkyl-O-(C=O)-, (C₃-C₁₀)cycloalkyl-O-(C=O)-,
(C₅-C₁₀)heterocyclic-O-(C=O)-, (C₁-C₆)alkyl-NH-(C=O)-,

- (C₃-C₁₀)cycloalkyl-NH-(C=O)-, phenyl-NH-(C=O)-,
(C₅-C₁₀)heterocyclic-NH-(C=O)-, (C₅-C₁₀)heteroaryl-NH-(C=O)-,
((C₁-C₆)alkyl)₂-N-(C=O)-, phenyl-[((C₁-C₆)alkyl)-N]-(C=O)-, hydroxy,
(C₁-C₆)alkoxy, perhalo(C₁-C₆)alkoxy, (C₃-C₁₀)cycloalkyl-O-, phenoxy,
5 (C₅-C₁₀)heterocyclic-O-, (C₅-C₁₀)heteroaryl-O-, (C₁-C₆)alkyl-(C=O)-O-,
(C₃-C₁₀)cycloalkyl-(C=O)-O-, phenyl-(C=O)-O-, (C₅-C₁₀)heterocyclic-(C=O)-O-,
(C₅-C₁₀)heteroaryl-(C=O)-O-, O₂N-, amino, (C₁-C₆)alkylamino,
((C₁-C₆)alkyl)₂-amino, formamidyl, (C₁-C₆)alkyl-(C=O)-NH-,
(C₃-C₁₀)cycloalkyl-(C=O)-NH-, phenyl-(C=O)-NH-,
10 (C₅-C₁₀)heterocyclic-(C=O)-NH-, (C₅-C₁₀)heteroaryl-(C=O)-NH-,
(C₁-C₆)alkyl-(C=O)-[((C₁-C₆)alkyl)-N]-, phenyl-(C=O)-[(C₁-C₆)alkyl-N]-,
(C₁-C₆)alkyl-SO₂NH-, (C₃-C₁₀)cycloalkyl-SO₂NH-, phenyl-SO₂NH-,
(C₅-C₁₀)heterocyclic-SO₂NH- and (C₅-C₁₀)heteroaryl-SO₂NH-;
wherein the phenyl or heteroaryl moiety of a R⁶ substituent is optionally
15 further substituted with at least one radical independently selected from the group
consisting of halo, (C₁-C₆)alkyl, (C₁-C₆)alkoxy, perfluoro(C₁-C₆)alkyl and
perfluoro(C₁-C₆)alkoxy,
with the proviso that R¹ contains at least one heteroatom.

- 20 2. A compound of claim 1, wherein R¹ is

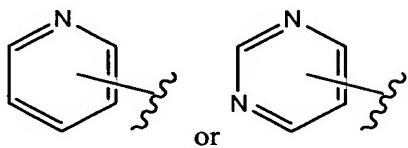


3. A compound of claim 1, wherein R¹ is



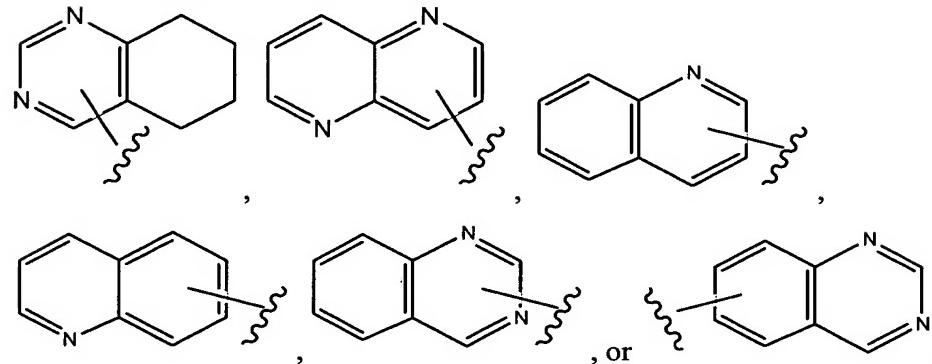
4. A compound of claim 1, wherein R¹ is

5

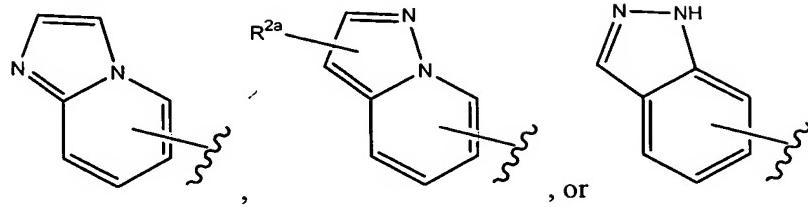


5. A compound of claim 1, wherein R¹ is

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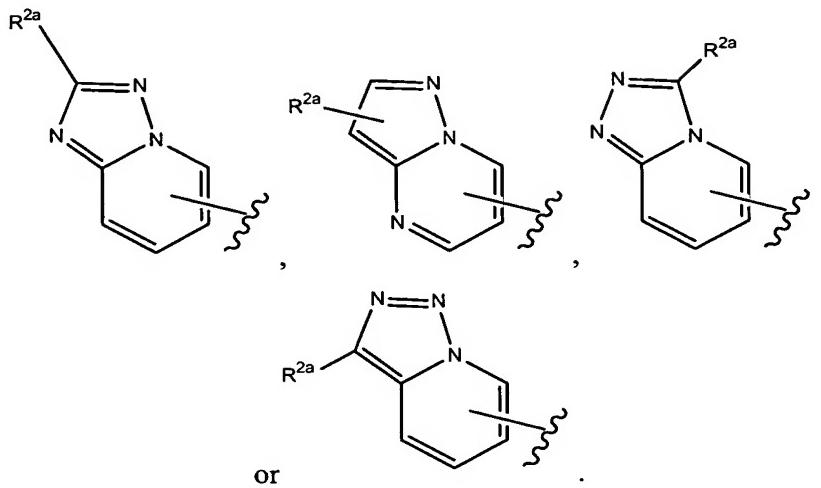


6. A compound of claim 1, wherein R¹ is



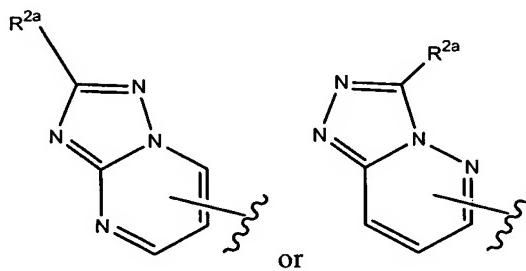
7. A compound of claim 1, wherein R¹ is

5



8. A compound of claim 1, wherein R¹ is

10



9. A compound of claim 1, wherein s is one to two; R³ is hydrogen or (C₁-C₆)alkyl; R⁴ is hydrogen, (C₁-C₆)alkyl, (C₃-C₁₀)cycloalkyl, amino, (C₁-C₆)alkylamino, (C₁-C₆)alkyl-(C=O)-, or (C₃-C₁₀)cycloalkyl-(C=O)-; and R⁶ is H or (C₁-C₆)alkyl.

10. A pharmaceutical composition comprising a compound of claim 1 and a pharmaceutically acceptable carrier.
- 5 11. A method of preventing or treating a TGF-related disease state in an animal or human comprising the step of administering a therapeutically effective amount of a compound of claim 1 to the animal or human suffering from the TGF-related disease state.
- 10 12. A method of claim 11, wherein said TGF-related disease state is selected from the group consisting of cancer, glomerulonephritis, diabetic nephropathy, hepatic fibrosis, pulmonary fibrosis, intimal hyperplasia and restenosis, scleroderma, and dermal scarring.